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(FILE 'HOME' ENTERED AT 15:07:14 ON 26 AUG 2005)

FILE 'HCAPLUS' ENTERED AT 15:07:22 ON 26 AUG 2005

1 US2005119219/PN OR (US2004-507472# OR WO2003-EP2538#)/AP,PRN L1

1 IT2002-PD64#/AP, PRN L2

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FILE 'HCAPLUS' ENTERED AT 15:10:03 ON 26 AUG 2005 1.3

1 L1, L2

FILE 'REGISTRY' ENTERED AT 15:10:13 ON 26 AUG 2005

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TRA L3 1- RN : 3 TERMS L4

FILE 'REGISTRY' ENTERED AT 15:10:15 ON 26 AUG 2005 3 SEA L4 L5

FILE 'WPIX' ENTERED AT 15:10:20 ON 26 AUG 2005 1 L3 L6

=> b hcap

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FILE COVERS 1907 - 26 Aug 2005 VOL 143 ISS 10 (20050825/ED) FILE LAST UPDATED: 25 Aug 2005

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This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d all 13 tot

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ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2005 ACS on STN
L_3
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- 2003:737790 HCAPLUS AN
- DN 139:262427
- ED Entered STN: 19 Sep 2003
- Ester derivatives of hyaluronic acid, preparation, hydrogel materials by ΤI photocuring, and biomedical use
- Bellini, Davide; Zanellato, Anna Maria IN
- Fidia Advanced Biopolymers S.R.L., Italy PΑ
- PCT Int. Appl., 27 pp. SO

CODEN: PIXXD2

- DT Patent
- LΑ English
- ICM C08B037-08 IC

ICS C08J003-28; C08J003-075

44-5 (Industrial Carbohydrates)

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Section cross-reference(s): 63
FAN.CNT 1
                                                                    DATE
     PATENT NO.
                         KIND DATE
                                            APPLICATION NO.
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                         A1
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             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
             PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT,
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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
             FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
             BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
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20050406 EP 2003-743875
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     EP 1519962
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                         A1
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                                 20050602 US 2003-507472
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                                 20020312 <--
PRAI IT 2002-PD64
                          Α
                         W
                                 20030312 <--
     WO 2003-EP2538
CLASS
 PATENT NO.
                CLASS PATENT FAMILY CLASSIFICATION CODES
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 WO 2003076475 ICM
                        C08B037-08
                        C08J003-28; C08J003-075
                 ICS
 WO 2003076475 ECLA A61L027/20+C08L5/08; A61L029/04D+C08L5/08;
                        A61L031/04D+C08L5/08; C08B037/00P2F
                                                                                <--
 US 2005119219 NCL
                        514/054.000
OS
     MARPAT 139:262427
     The hyaluronic acid ester derivs., have carboxylic groups are partially
AB
     esterified with hydroxy groups of propiophenone derivs. Thus, 6.21 g
     tetrabutylammonium salt of hyaluronic acid, mol. weight 180,000 Da (10 meq.)
     are solubilized in 248 mL DMSO at room temperature, 2 g 2-hydroxy-4-(2-
     hydroxyethoxy)-2-methylpropiophenone (HHMP) bromide (7 meq.) are added,
     the solution is maintained at 37° for 48 h, 2.5% NaCl solution is added
     and the mixture is poured under stirring into 750 mL acetone, precipitating,
     filtering and washing three times with 100 mL acetone:water 5:1, three
     times with 100 mL acetone, and lastly vacuum-drying 24 h at 30°
     gave 5.3 g HHMP ester product.
     hyaluronic acid ester prepn photocuring hydrogel biol use
st
IT
     Drug delivery systems
        (controlled-release; hyaluronic acid esters photocured for hydrogels
        for)
IT
     Cell proliferation
        (fibroblast, scaffolds; hyaluronic acid esters photocured for hydrogels
        for)
IT
     Medical goods
        (hyaluronic acid esters photocured for hydrogels for)
IT
     Drug delivery systems
        (hydrogels; hyaluronic acid esters photocured for hydrogels for)
IT
     Fibroblast
        (proliferation, scaffolds; hyaluronic acid esters photocured for
        hydrogels for)
     124-22-1DP, Dodecylamine, amide with hyaluronic acid esters
IT
     601468-77-3DP, amide with dodecylamine 601468-77-3P, Hyaluronic acid
     2-hydroxy-4-(2-hydroxyethoxy)-2-methylpropiophenone ester sodium salt
     601468-78-4P
     RL: BUU (Biological use, unclassified); IMF (Industrial manufacture); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
        (hyaluronic acid esters photocured for hydrogels)
              THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 6
RE
(1) Aeschlimann; WO 0016818 A 2000 HCAPLUS
```

- (2) Fidia Advanced Biopolymers Srl; WO 9637519 A 1996 HCAPLUS
- (3) Hercules Incorporated; EP 0749982 A 1996 HCAPLUS
- (4) Nguyen, K; BIOMATERIALS 2002, V23(22), P4307 HCAPLUS
- (5) Seikagaku Corporation; WO 9718244 A 1997 HCAPLUS
- (6) Waki; US 6031017 A 2000 HCAPLUS

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YOU HAVE REQUESTED DATA FROM FILE 'HCAPLUS' - CONTINUE? (Y) /N:n

=> d ide 15 tot

- L5 ANSWER 1 OF 3 REGISTRY COPYRIGHT 2005 ACS on STN
- RN 601468-78-4 REGISTRY
- ED Entered STN: 09 Oct 2003
- CN Hyaluronic acid, ester with 2-hydroxy-1-[4-(2-hydroxyethoxy)phenyl]-2-methyl-1-propanone, phenylmethyl ester, sodium salt (9CI) (CA INDEX NAME)
- MF C12 H16 O4 . \times C7 H8 O . \times Na . \times Unspecified
- PCT Manual registration, Polyester, Polyester formed
- SR C
- LC STN Files: CA, CAPLUS, USPATFULL

CM 1

CRN 106797-53-9 CMF C12 H16 O4

CM 2

CRN 9004-61-9 CMF Unspecified CCI PMS, MAN

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

CM 3

CRN 100-51-6 CMF C7 H8 O

HO- CH2- Ph

1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L5 ANSWER 2 OF 3 REGISTRY COPYRIGHT 2005 ACS on STN

RN 601468-77-3 REGISTRY

ED Entered STN: 09 Oct 2003

CN Hyaluronic acid, ester with 2-hydroxy-1-[4-(2-hydroxyethoxy)phenyl]-2methyl-1-propanone, sodium salt (9CI) (CA INDEX NAME)
OTHER NAMES:

CN Hyaluronic acid 2-hydroxy-4-(2-hydroxyethoxy)-2-methylpropiophenone ester sodium salt

MF $\,$ C12 H16 O4 $\,$ x Na $\,$ x Unspecified

PCT Manual registration, Polyester, Polyester formed

SR C

LC STN Files: CA, CAPLUS, USPATFULL

CM 1

CRN 106797-53-9 CMF C12 H16 O4

CM 2

CRN 9004-61-9 CMF Unspecified CCI PMS, MAN

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*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
               1 REFERENCES IN FILE CA (1907 TO DATE)
                1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
                1 REFERENCES IN FILE CAPLUS (1907 TO DATE)
     ANSWER 3 OF 3 REGISTRY COPYRIGHT 2005 ACS on STN
L_5
     124-22-1 REGISTRY
RN
ED
     Entered STN: 16 Nov 1984
     1-Dodecanamine (9CI) (CA INDEX NAME)
CN
OTHER CA INDEX NAMES:
   Dodecylamine (8CI)
OTHER NAMES:
    1-Aminododecane
CN
     1-Dodecylamine
CN
CN
    Alamine 4
CN
    Amine BB
CN
    Armeen 12D
CN
    Farmin 20D
CN
     Kemamine P 690
CN
    Lauramine
CN
    Laurinamine
CN
    Laurylamine
CN
    Monododecylamine
CN
    Monolaurylamine
CN
    n-Dodecylamine
    n-Laurylamine
CN
    Nissan Amine BB
CN
     3D CONCORD
FS
DR
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MF
     C12 H27 N
CI
     COM
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LC
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       ENCOMPLIT, ENCOMPLIT2, ENCOMPPAT, ENCOMPPAT2, GMELIN*, HODOC*, HSDB*,
       IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MSDS-OHS, NIOSHTIC, PIRA, PROMT,
       PS, RTECS*, SPECINFO, SYNTHLINE, TOXCENTER, TULSA, ULIDAT, USPAT2,
       USPATFULL, VTB
         (*File contains numerically searchable property data)
     Other Sources: DSL**, EINECS**, TSCA**
         (**Enter CHEMLIST File for up-to-date regulatory information)
H_2N^- (CH<sub>2</sub>)<sub>11</sub>-Me
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
            4960 REFERENCES IN FILE CA (1907 TO DATE)
             744 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            4968 REFERENCES IN FILE CAPLUS (1907 TO DATE)
              15 REFERENCES IN FILE CAOLD (PRIOR TO 1967)
=> b wpix;d all drn dcn 16 tot
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23 AUG 2005

200554 DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

FILE LAST UPDATED:

MOST RECENT DERWENT UPDATE:

<20050823/UP> <200554/DW>

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    DOCUMENTATION NOW AVAILABLE IN DERWENT WORLD PATENTS INDEX
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    FOR FURTHER DETAILS: http://www.thomsonderwent.com/dwpifv <<<
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    ANSWER 1 OF 1 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
L6
     2003-876823 [81]
AN
                       WPIX
DNC
    C2003-247398
     Ester derivatives of hyaluronic acid or hyaluronic acid derivatives, for
TI
    use as hydrogel material, comprising some carboxylic groups which are
     esterified with propiophenone derivatives.
DC
    A11 A96 B05 B07 D22
    BELLINI, D; ZANELLATO, A M
IN
     (FIDI-N) FIDIA ADVANCED BIOPOLYMERS SRL; (BELL-I) BELLINI D; (ZANE-I)
     ZANELLATO A M
CYC
    104
PΙ
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                                                27
                                                      C08B037-08
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           DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR
           KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NI NO NZ OM PH PL
           PT RO RU SC SD SE SG SK SL TJ TM TN TR TT TZ UA UG US UZ VC VN YU
           ZA ZM ZW
    AU 2003227050
                    A1 20030922 (200431)
                                                      C08B037-08
                    A1 20050406 (200523) EN
                                                      C08B037-08
    EP 1519962
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    US 2005119219
                   A1 20050602 (200537)
                                                      A61K031-715
    WO 2003076475 A1 WO 2003-EP2538 20030312; AU 2003227050 A1 AU
ADT
     2003-227050 20030312; EP 1519962 A1 EP 2003-743875 20030312, WO
     2003-EP2538 20030312; US 2005119219 A1 WO 2003-EP2538
     20030312, US 2004-507472 20040910
FDT AU 2003227050 A1 Based on WO 2003076475; EP 1519962 A1 Based on WO
     2003076475
PRAI IT 2002-PD64
                          20020312
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     ICS A01N043-04; C08J003-075; C08J003-28
AB
     WO2003076475 A UPAB: 20031216
    NOVELTY - Ester derivatives of hyaluronic acid or its derivatives,
     comprise some carboxylic groups which are esterified with propiophenone
     derivatives (I).
         DETAILED DESCRIPTION - Ester derivatives of hyaluronic acid or its
     derivatives, comprise some carboxylic groups which are esterified with
    propiophenone derivatives of formula (I).
          R = hydroxy, or 1-20C alkyloxy or heterocycle (both substituted by
    hydroxy group(s)); and
          R1-R3 = H, hydroxy, or 1-20C alkyl or 1-20C alkyloxy (optionally
     substituted with hydroxy group(s)).
```

INDEPENDENT CLAIMS are also included for:

- (a) preparation of the ester derivatives, comprising reacting hyaluronic acid or its derivatives with the bromide of (I), where hydroxy group(s) of the substituent R is replaced by Bromine;
- (b) a hydrogel material consisting of a cross-linked product obtained by photocuring an ester derivative; and
- (c) a kit for implanting engineered cartilage by arthroscopic surgery, comprising an ester derivative dissolved in water or in an aqueous solution, a container for the ester derivative, and an endoscopic probe with optic fibers for the photocuring of the ester derivative.

USE - Useful in hydrogel material for biomedical materials, healthcare products, surgical articles consisting of catheters, guide channels, cardiac valves, vascular stents, soft tissue prostheses, prostheses of animal origin such as porcine heart valves, artificial tendons and organs, contact lens, and intra-ocular lenses, blood oxygenators, blood bags, surgical instruments, filtrations systems and laboratory instruments, for scaffolds for the growth of human and animal, pharmaceutical composition for topical, subcutaneous, intramuscular, intra-articular and intra-medullar administration, in the prevention of surgical adhesions, for the preparation of engineered connective tissues, engineered cartilage, viscoelastic substitutes of the nucleus pulpous of the intervertebral disk and visco-integrators of the vitreous humor (claimed).

ADVANTAGE - The hydrogel materials show valuable properties, and have absence of C=C unsaturation in the ester derivatives, without the addition of any component acting as catalyst for the cross-linking reaction and without any unsaturation within the molecule. The hydrogel materials also show sterility, excellent viscoelastic properties, compact, wall-to-wall type, three-dimensional structure, greater mechanical resistance, and have viscoelastic properties that vary according to how long they are exposed to irradiation, and to the type of aqueous solution used to obtain the hydrogel.

```
to irradiation, and to the type of aqueous solution used to obtain the
     hydrogel.
     Dwg.0/1
FS
     CPI
     AB; GI; DCN
FA
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MC
            B04-L01; B04-N04; B06-H; B07-H; B10-E02; B10-E04; B10-F02; B12-M10;
            B14-E11; D09-C01
     0272-S; 0272-U; 0274-S; 0274-U; 0714-P; 0714-S; 0714-U; 1064-P; 1064-S;
      1064-U; 1706-S; 1706-U
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          R06437-T; R06437-Q; R06437-M; R06437-P
*02* DCN: RA0121-K; RA0121-T; RA0121-Q; RA0121-M; RA0121-P
          *03* DCN: RA0019-K; RA0019-T; RA0019-Q; RA0019-M; RA0019-P
          *04* DCN: RA0120-K; RA0120-T; RA0120-Q; RA0120-M; RA0120-P
          *13* DCN: RA00H3-K; RA00H3-T; RA00H3-Q; RA00H3-M; RA00H3-P
          *14* DCN: RA00C8-K; RA00C8-T; RA00C8-Q; RA00C8-M; RA00C8-P
*15* DCN: RA00GC-K; RA00GC-T; RA00GC-Q; RA00GC-M; RA00GC-P
*05* DCN: RABXF5-K; RABXF5-T; RABXF5-Q; RABXF5-M; RABXF5-P
     M2
          *06* DCN: RABXF6-K; RABXF6-T; RABXF6-Q; RABXF6-M; RABXF6-P
     M2
          *07* DCN: RABXF7-K; RABXF7-T; RABXF7-Q; RABXF7-M; RABXF7-P
          *08* DCN: R03586-K; R03586-T; R03586-Q; R03586-M; R03586-P
     M2
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*10* DCN: R00714-K; R00714-T; R00714-Q; R00714-M; R00714-P
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                      0109-97801-P
          *16* DCN: R00274-K; R00274-V; R00274-U
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*18* DCN: R00272-K; R00272-V; R00272-U
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=> b home FILE 'HOME' ENTERED AT 15:12:00 ON 26 AUG 2005

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